

A-378CIP5 rev.ST25.txt SEQUENCE LISTING

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CHANG, MING-SHI

SENALDI, GIORGIO

RECEIVED

OCT 0 8 2002

TECH CENTER 1600/2900

<120> COMBINATION THERAPY FOR CONDITIONS LEADING TO BONE LOSS

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<141> 2000-07-10

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aca atg aac aag tgg ctg tgc tgt gca ctc ctg gtg ttc ttg gac atc Met Asn Lys Trp Leu Cys Cys Ala Leu Leu Val Phe Leu Asp Ile 1 5 10 15	168						
att gaa tgg aca acc cag gaa acc ttt cct cca aaa tac ttg cat tat Ile Glu Trp Thr Thr Gln Glu Thr Phe Pro Pro Lys Tyr Leu His Tyr 20 25 30	216						
gac cca gaa acc gga cgt cag ctc ttg tgt gac aaa tgt gct cct ggc Asp Pro Glu Thr Gly Arg Gln Leu Leu Cys Asp Lys Cys Ala Pro Gly 35 40 45	264						
acc tac cta aaa cag cac tgc aca gtc agg agg aag aca ctg tgt gtc Thr Tyr Leu Lys Gln His Cys Thr Val Arg Arg Lys Thr Leu Cys Val 50 55 60	312						
cct tgc cct gac tac tct tat aca gac agc tgg cac acg agt gat gaa Pro Cys Pro Asp Tyr Ser Tyr Thr Asp Ser Trp His Thr Ser Asp Glu 65 70 75	360						
tgc gtg tac tgc agc ccc gtg tgc aag gaa ctg cag acc gtg aaa cag Cys Val Tyr Cys Ser Pro Val Cys Lys Glu Leu Gln Thr Val Lys Gln 80 85 90 95	408						
gag tgc aac cgc acc cac aac cga gtg tgc gaa tgt gag gaa ggg cgc Glu Cys Asn Arg Thr His Asn Arg Val Cys Glu Cys Glu Glu Gly Arg 100 105 110	456						
tac ctg gag ctc gaa ttc tgc ttg aag cac cgg agc tgt ccc cca ggc Tyr Leu Glu Leu Glu Phe Cys Leu Lys His Arg Ser Cys Pro Pro Gly 115 120 125	504						
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agg ttt gct gtg cct acc aag att ata ccg aat tgg ctg agt gtt ctg Arg Phe Ala Val Pro Thr Lys Ile Ile Pro Asn Trp Leu Ser Val Leu 210 215 220	792						
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ata aaa cgg aga cac agc tcg caa gag caa act ttc cag cta ctt aag Ile Lys Arg Arg His Ser Ser Gln Glu Gln Thr Phe Gln Leu Leu Lys 240 245 250 255 Page 34	888						

ctg tgg aag cat caa aac aga gac cag gaa atg gtg aag aag atc atc Leu Trp Lys His Gln Asn Arg Asp Gln Glu Met Val Lys Lys Ile Ile 260 265 270	936
caa gac att gac ctc tgt gaa agc agt gtg caa cgg cat atc ggc cac Gln Asp Ile Asp Leu Cys Glu Ser Ser Val Gln Arg His Ile Gly His 275 280 285	984
gcg aac ctc acc aca gag cag ctc cgc atc ttg atg gag agc ttg cct Ala Asn Leu Thr Thr Glu Gln Leu Arg Ile Leu Met Glu Ser Leu Pro 290 295 300	1032
ggg aag aag atc agc cca gac gag att gag aga acg aga aag acc tgc Gly Lys Lys Ile Ser Pro Asp Glu Ile Glu Arg Thr Arg Lys Thr Cys 305 310 315	1080
aaa ccc agc gag cag ctc ctg aag cta ctg agc ttg tgg agg atc aaa Lys Pro Ser Glu Gln Leu Leu Lys Leu Leu Ser Leu Trp Arg Ile Lys 320 335 330 335	1128
aat gga gac caa gac acc ttg aag ggc ctg atg tac gca ctc aag cac Asn Gly Asp Gln Asp Thr Leu Lys Gly Leu Met Tyr Ala Leu Lys His 340 345 350	1176
ttg aaa gca tac cac ttt ccc aaa acc gtc acc cac agt ctg agg aag Leu Lys Ala Tyr His Phe Pro Lys Thr Val Thr His Ser Leu Arg Lys 355 360 365	1224
acc atc agg ttc ttg cac agc ttc acc atg tac cga ttg tat cag aaa Thr Ile Arg Phe Leu His Ser Phe Thr Met Tyr Arg Leu Tyr Gln Lys 370 375 380	1272
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Glu Trp Thr	Thr Gln G	lu Thr Phe	Pro Pro Lys	Tyr Leu His	Tyr Asp		
Pro Glu Thr 35	Gly Arg G	iln Leu Leu 40	Cys Asp Lys	Cys Ala Pro 45	Gly Thr		
Tyr Leu Lys 50	Gln His C	Cys Thr Val	Arg Arg Lys	Thr Leu Cys	Val Pro		
		1					
65 Pro ASP	Tyr Ser I 7	'O	Ser Trp His 75	inr Ser Asp	80		
val Tyr Cys	Ser Pro V 85	al Cys Lys	Glu Leu Gln 90	Thr Val Lys	Gln Glu 95		
Cys Asn Arg	Thr His A 100	Asn Arg Val	Cys Glu Cys 105	Glu Glu Gly 110	Arg Tyr		
			His Arg Ser		Gly Leu		
115		120		125			
Gly Val Leu 130	Gln Ala G	oly Thr Pro 135	Glu Arg Asn	Thr Val Cys 140	Lys Arg		
CVS Pro Asn	Gly Phe B	he Ser Glv	Glu Thr Ser	Ser Lys Ala	Pro Cvs		
145		150	155	Jer Lys Ala	160		
Arg Lys His	Thr Asn C	ys Ser Ser	Leu Gly Leu 170	Leu Leu Ile	Gln Lys 175		
	103		1,0		<u> </u>		
Gly Asn Ala	Thr His A 180	Asp Asn Val	Cys Ser Gly 185	190	Ala Thr		
Page 36							

Gln Asn Cys Glu Ile Asp Val Thr Leu Cys Glu Glu Ala Phe Phe Arg 195 200 205 Phe Ala Val Pro Thr Lys Ile Ile Pro Asn Trp Leu Ser Val Leu Val 210 215 220 Asp Ser Leu Pro Gly Thr Lys Val Asn Ala Glu Ser Val Glu Arg Ile 225 230 235 240 Lys Arg Arg His Ser Ser Gln Glu Gln Thr Phe Gln Leu Leu Lys Leu 245 250 255 Trp Lys His Gln Asn Arg Asp Gln Glu Met Val Lys Lys Ile Ile Gln
260 265 270 Asp Ile Asp Leu Cys Glu Ser Ser Val Gln Arg His Ile Gly His Ala 275 280 285 Asn Leu Thr Thr Glu Gln Leu Arg Ile Leu Met Glu Ser Leu Pro Gly 290 295 300 Lys Lys Ile Ser Pro Asp Glu Ile Glu Arg Thr Arg Lys Thr Cys Lys 305 310 315 320Pro Ser Glu Gln Leu Leu Lys Leu Leu Ser Leu Trp Arg Ile Lys Asn 325 330 335 Gly Asp Gln Asp Thr Leu Lys Gly Leu Met Tyr Ala Leu Lys His Leu 340 345 350 Lys Ala Tyr His Phe Pro Lys Thr Val Thr His Ser Leu Arg Lys Thr 355 360 365 Ile Arg Phe Leu His Ser Phe Thr Met Tyr Arg Leu Tyr Gln Lys Leu 370 380 Phe Leu Glu Met Ile Gly Asn Gln Val Gln Ser Val Lys Ile Ser Cys 385 390 395 400

Leu

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<211> 1325

<212> DNA

<213> Mus musculus

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Page 38

Phe Gl 17		Leu	Leu	Ile	Gln 175	A- Lys	378C Gly	IP5 Asn	rev. Ala	ST25 Thr 180	.txt His	Asp	Asn	Cys	
tgt to Cys Se 185	c gga r Gly	aac Asn	aga Arg	gaa Glu 190	gcc Ala	acg Thr	caa Gln	aag Lys	tgt Cys 195	gga Gly	ata Ile	gat Asp	gtc Val	acc Thr 200	690
ctg tg Leu Cy															738
cca aa Pro As	t tgg n Trp	ctg Leu 220	agt Ser	gtt Val	ttg Leu	gtg Val	gac Asp 225	agt Ser	ttg Leu	cct Pro	ggg Gly	acc Thr 230	aaa Lys	gtg Val	786
aat gc Asn Al															834
caa ac Gln Th 25	r Phe														882
gaa at Glu Me 265															930
gtg ca Val Gl	g cgg n Arg	cat His	ctc Leu 285	ggc Gly	cac His	tcg Ser	aac Asn	ctc Leu 290	acc Thr	aca Thr	gag Glu	cag Gln	ctt Leu 295	ctt Leu	978
gcc tt Ala Le	g atg u Met	gag G1u 300	agc Ser	ctg Leu	cct Pro	ggg Gly	aag Lys 305	aag Lys	atc Ile	agc Ser	cca Pro	gaa Glu 310	gag Glu	att Ile	1026
gag ag Glu Ar															1074
ctc ag Leu Se 33	r Leu	tgg Trp	agg Arg	atc Ile	aaa Lys 335	aat Asn	ggt Gly	gac Asp	caa Gln	gac Asp 340	acc Thr	ttg Leu	aag Lys	ggc Gly	1122
ctg at Leu Me 345															1170
gtc ac Val Th	c cac r His	agt Ser	ctg Leu 365	agg Arg	aag Lys	acc Thr	atg Met	agg Arg 370	ttc Phe	ctg Leu	cac His	agc Ser	ttc Phe 375	aca Thr	1218
atg ta Met Ty	c aga r Arg	ctg Leu 380	tat Tyr	cag Gln	aag Lys	ctc Leu	ttt Phe 385	tta Leu	gaa Glu	atg Met	ata Ile	ggg GTy 390	aat Asn	cag Gln	1266
gtt ca Val Gl	a tcc n Ser 395	gtg Val	aaa Lys	ata Ile	agc Ser	tgc Cys 400	tta Leu	taad	tago	gaa t	ggto	acto	99		1313
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<221> misc_feature

<222> (11)..(11)

<223> At position 11, R is a purine.

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Pro Glu Thr Gly His Gln Leu Leu Cys Asp Lys Cys Ala Pro Gly Thr 35 40 45

Tyr Leu Lys Gln His Cys Thr Val Arg Arg Lys Thr Leu Cys Val Pro 50 60

Cys Pro Asp His Ser Tyr Thr Asp Ser Trp His Thr Ser Asp Glu Cys 65 70 75 80

Val Tyr Cys Ser Pro Val Cys Lys Glu Leu Gln Ser Val Lys Gln Glu 85 90 95

Cys Asn Arg Thr His Asn Arg Val Cys Glu Cys Glu Glu Gly Arg Tyr 100 105 110

Leu Glu Ile Glu Phe Cys Leu Lys His Arg Ser Cys Pro Pro Gly Ser 115 120 125

Gly Val Val Gln Ala Gly Thr Pro Glu Arg Asn Thr Val Cys Lys 130 135 140

Cys Pro Asp Gly Phe Phe Ser Gly Glu Thr Ser Ser Lys Ala Pro Cys 145 150 155 160

Ile Lys His Thr Asn Cys Ser Thr Phe Gly Leu Leu Leu Ile Gln Lys 165 170 175

Gly Asn Ala Thr His Asp Asn Cys Cys Ser Gly Asn Arg Glu Ala Thr 180 185 190

Gln Lys Cys Gly Ile Asp Val Thr Leu Cys Glu Glu Ala Phe Phe Arg 195 200 205

Phe Ala Val Pro Thr Lys Ile Ile Pro Asn Trp Leu Ser Val Leu Val 210 220 Page 40

Asp Ser Leu Pro Gly Thr Lys Val Asn Ala Glu Ser Val Glu Arg Ile 225 230 235 240 Lys Arg Arg His Ser Ser Gln Glu Gln Thr Phe Gln Leu Leu Lys Leu 245 250 255 Trp Lys His Gln Asn Arg Asp Gln Glu Met Val Lys Lys Ile Ile Gln 260 265 270 Asp Ile Asp Leu Cys Glu Ser Ser Val Gln Arg His Leu Gly His Ser 275 280 285 Asn Leu Thr Thr Glu Gln Leu Leu Ala Leu Met Glu Ser Leu Pro Gly 290 295 300 Lys Lys Ile Ser Pro Glu Glu Ile Glu Arg Thr Arg Lys Thr Cys Lys 305 310 315 Ser Ser Glu Gln Leu Leu Lys Leu Leu Ser Leu Trp Arg Ile Lys Asn 325 330 335 Gly Asp Gln Asp Thr Leu Lys Gly Leu Met Tyr Ala Leu Lys His Leu 340 345 350 Lys Thr Ser His Phe Pro Lys Thr Val Thr His Ser Leu Arg Lys Thr 355 360 365 Met Arg Phe Leu His Ser Phe Thr Met Tyr Arg Leu Tyr Gln Lys Leu 370 375 380 Phe Leu Glu Met Ile Gly Asn Gln Val Gln Ser Val Lys Ile Ser Cys 385 390 395 400 Leu

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<213> Homo sapiens

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<223>

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<222>	(63)(63)
<223>	At position 63, Y is a pyrimidine.

<400> 124 gtatatataa cgtgatgagc gtacgggtgc ggagacgcac cggcgcgctc gcccagccgc	60
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gcg ctc gtg ttt ctg gac atc tcc att aag tgg acc acc cag gaa acg Ala Leu Val Phe Leu Asp Ile Ser Ile Lys Trp Thr Thr Gln Glu Thr 10 15 20	163
ttt cct cca aag tac ctt cat tat gac gaa gaa acc tct cat cag ctg Phe Pro Pro Lys Tyr Leu His Tyr Asp Glu Glu Thr Ser His Gln Leu 25 30 35	211
ttg tgt gac aaa tgt cct cct ggt acc tac cta aaa caa cac tgt aca Leu Cys Asp Lys Cys Pro Pro Gly Thr Tyr Leu Lys Gln His Cys Thr 40 45 50 55	259
gca aag tgg aag tcc gtg tgc gcc cct tgc cct gac cac tac tac aca Ala Lys Trp Lys Ser Val Cys Ala Pro Cys Pro Asp His Tyr Tyr Thr 60 65 70	307
gac agc tgg cac acc agt gac gag tgt cta tac tgc agc ccc gtg tgc Asp Ser Trp His Thr Ser Asp Glu Cys Leu Tyr Cys Ser Pro Val Cys 75 80 85	355
aag gag ctg cag tac gtc aag cag gag tgc aat cgc acc cac aac cgc Lys Glu Leu Gln Tyr Val Lys Gln Glu Cys Asn Arg Thr His Asn Arg 90 95 100	403
gtg tgc gaa tgc aag gaa ggg cgc tac ctt gag ata gag ttc tgc ttg Val Cys Glu Cys Lys Glu Gly Arg Tyr Leu Glu Ile Glu Phe Cys Leu 105 110 115	451
aaa cat agg agc tgc cct cct gga ttt gga gtg gtg caa gct gga acc Lys His Arg Ser Cys Pro Pro Gly Phe Gly Val Val Gln Ala Gly Thr 120 125 130 135	499
cca gag cga aat aca gtt tgc aaa aga tgt cca gat ggg ttc ttc tca Pro Glu Arg Asn Thr Val Cys Lys Arg Cys Pro Asp Gly Phe Phe Ser 140 145 150	547
aat gag acg tca tct aaa gca ccc tgt aga aaa cac aca aat tgc agt Asn Glu Thr Ser Ser Lys Ala Pro Cys Arg Lys His Thr Asn Cys Ser 155 160 165	595
gtc ttt ggt ctc ctg cta act cag aaa gga aat gca aca cac gac aac Val Phe Gly Leu Leu Thr Gln Lys Gly Asn Ala Thr His Asp Asn 170 175 180	643
ata tgt tcc gga aac agt gaa tca act caa aaa tgt gga ata gat gtt Ile Cys Ser Gly Asn Ser Glu Ser Thr Gln Lys Cys Gly Ile Asp Val 185 190 195	691
acc ctg tgt gag gag gca ttc ttc agg ttt gct gtt cct aca aag ttt Page 42	739

A-378CIP5 rev.ST25.txt Thr Leu Cys Glu Glu Ala Phe Phe Arg Phe Ala Val Pro Thr Lys Phe 200 205 210 215	
acg cct aac tgg ctt agt gtc ttg gta gac aat ttg cct ggc acc aaa Thr Pro Asn Trp Leu Ser Val Leu Val Asp Asn Leu Pro Gly Thr Lys 220 225 230	787
gta aac gca gag agt gta gag agg ata aaa cgg caa cac agc tca caa Val Asn Ala Glu Ser Val Glu Arg Ile Lys Arg Gln His Ser Ser Gln 235 240 245	835
gaa cag act ttc cag ctg ctg aag tta tgg aaa cat caa aac aaa gcc Glu Gln Thr Phe Gln Leu Leu Lys Leu Trp Lys His Gln Asn Lys Ala 250 255 260	883
caa gat ata gtc aag aag atc atc caa gat att gac ctc tgt gaa aac Gln Asp Ile Val Lys Lys Ile Ile Gln Asp Ile Asp Leu Cys Glu Asn 265 270 275	931
agc gtg cag cgg cac att gga cat gct aac ctc acc ttc gag cag ctt Ser Val Gln Arg His Ile Gly His Ala Asn Leu Thr Phe Glu Gln Leu 280 285 290 295	979
cgt agc ttg atg gaa agc tta ccg gga aag aaa gtg gga gca gaa gac Arg Ser Leu Met Glu Ser Leu Pro Gly Lys Lys Val Gly Ala Glu Asp 300 305 310	1027
att gaa aaa aca ata aag gca tgc aaa ccc agt gac cag atc ctg aag Ile Glu Lys Thr Ile Lys Ala Cys Lys Pro Ser Asp Gln Ile Leu Lys 315 320 325	1075
ctg ctc agt ttg tgg cga ata aaa aat ggc gac caa gac acc ttg aag Leu Leu Ser Leu Trp Arg Ile Lys Asn Gly Asp Gln Asp Thr Leu Lys 330 335 340	1123
ggc cta atg cac gca cta aag cac tca aag acg tac cac ttt ccc aaa Gly Leu Met His Ala Leu Lys His Ser Lys Thr Tyr His Phe Pro Lys 345 350 355	1171
act gtc act cag agt cta aag aag acc atc agg ttc ctt cac agc ttc Thr Val Thr Gln Ser Leu Lys Lys Thr Ile Arg Phe Leu His Ser Phe 360 365 370 375	1219
aca atg tac aaa ttg tat cag aag tta ttt tta gaa atg ata ggt aac Thr Met Tyr Lys Leu Tyr Gln Lys Leu Phe Leu Glu Met Ile Gly Asn 380 385 390	1267
cag gtc caa tca gta aaa ata agc tgc tta taactggaaa tggccattga Gln Val Gln Ser Val Lys Ile Ser Cys Leu 395 400	1317
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<220>	
<221> misc_feature	

<222> (63)..(63)

<223> At position 63, Y is a pyrimidine.

<400> 125

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Glu Glu Thr Ser His Gln Leu Leu Cys Asp Lys Cys Pro Pro Gly Thr 35 40 45

Tyr Leu Lys Gln His Cys Thr Ala Lys Trp Lys Ser Val Cys Ala Pro 50 55 60

Cys Pro Asp His Tyr Tyr Thr Asp Ser Trp His Thr Ser Asp Glu Cys 65 70 75 80

Leu Tyr Cys Ser Pro Val Cys Lys Glu Leu Gln Tyr Val Lys Gln Glu 85 90 95

Cys Asn Arg Thr His Asn Arg Val Cys Glu Cys Lys Glu Gly Arg Tyr 100 105 110

Leu Glu Ile Glu Phe Cys Leu Lys His Arg Ser Cys Pro Pro Gly Phe 115 120 125

Gly Val Val Gln Ala Gly Thr Pro Glu Arg Asn Thr Val Cys Lys Arg 130 135 140

Cys Pro Asp Gly Phe Phe Ser Asn Glu Thr Ser Ser Lys Ala Pro Cys 145 150 155 160

Arg Lys His Thr Asn Cys Ser Val Phe Gly Leu Leu Leu Thr Gln Lys 165 170 175

Gly Asn Ala Thr His Asp Asn Ile Cys Ser Gly Asn Ser Glu Ser Thr 180 185 190

Gln Lys Cys Gly Ile Asp Val Thr Leu Cys Glu Glu Ala Phe Phe Arg 195 200 205

Phe Ala Val Pro Thr Lys Phe Thr Pro Asn Trp Leu Ser Val Leu Val 210 215 220

Asp Asn Leu Pro Gly Thr Lys Val Asn Ala Glu Ser Val Glu Arg Ile 225 230 235 240

Lys Arg Gln His Ser Ser Gln Glu Gln Thr Phe Gln Leu Leu Lys Leu 245 250 255 Page 44

Trp Lys His Gln Asn Lys Ala Gln Asp Ile Val Lys Lys Ile Ile Gln 260 265 270

Asp Ile Asp Leu Cys Glu Asn Ser Val Gln Arg His Ile Gly His Ala 275 280 285

Asn Leu Thr Phe Glu Gln Leu Arg Ser Leu Met Glu Ser Leu Pro Gly 290 295 300

Lys Lys Val Gly Ala Glu Asp Ile Glu Lys Thr Ile Lys Ala Cys Lys 305 310 315 320

Pro Ser Asp Gln Ile Leu Lys Leu Leu Ser Leu Trp Arg Ile Lys Asn 325 330 335

Gly Asp Gln Asp Thr Leu Lys Gly Leu Met His Ala Leu Lys His Ser 340 345 350

Lys Thr Tyr His Phe Pro Lys Thr Val Thr Gln Ser Leu Lys Lys Thr 355 360 365

Ile Arg Phe Leu His Ser Phe Thr Met Tyr Lys Leu Tyr Gln Lys Leu 370 380

Phe Leu Glu Met Ile Gly Asn Gln Val Gln Ser Val Lys Ile Ser Cys 385 390 395 400

Leu

<210> 126

<211> 139

<212> PRT

<213> Homo sapiens

<400> 126

Cys Pro Gln Gly Lys Tyr Ile His Pro Gln Asn Asn Ser Ile Cys Cys
1 10 15

Thr Lys Cys His Lys Gly Thr Tyr Leu Tyr Asn Asp Cys Pro Gly Pro 20 25 30

Gly Gln Asp Thr Asp Cys Arg Glu Cys Glu Ser Gly Ser Phe Thr Ala 35 40 45

Ser Glu Asn His Leu Arg His Cys Leu Ser Cys Ser Lys Cys Arg Lys 50 60 Page 45

Glu Met Gly Gln Val Glu Ile Ser Ser Cys Thr Val Asp Arg Asp Thr 65 70 75 80

Val Cys Gly Cys Arg Lys Asn Gln Tyr Arg His Tyr Trp Ser Glu Asn 85 90 95

Leu Phe Gln Cys Phe Asn Cys Ser Leu Cys Leu Asn Gly Thr Val His 100 105 110

Leu Ser Cys Gln Glu Lys Gln Asn Thr Val Cys Thr Cys His Ala Gly 115 120 125

Phe Phe Leu Arg Glu Asn Glu Cys Val Ser Cys 130 135

<210> 127

<211> 48

<212> DNA

<213> Artificial Sequence

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<223> hu0PG

<400> 127

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<210> 128

<211> 219

<212> PRT

<213> Rattus rattus

<400> 128

Met Leu Gly Ile Trp Thr Leu Leu Pro Leu Val Leu Thr Ser Val Ala 1 5 10 15

Arg Leu Ser Ser Lys Ser Val Asn Ala Gln Val Thr Asp Ile Asn Ser 20 25 30

Lys Gly Leu Glu Leu Arg Lys Thr Val Thr Thr Val Glu Thr Gln Asn 35 40 45

Leu Glu Gly Leu His His Asp Gly Gln Phe Cys His Lys Pro Cys Pro 50 55 60

48

A-378CIPS rev. ST25.txt Asp Glu Pro Room Asp Cys Thr Val Asn Gly Asp Glu Pro Room Asp Cys Val Pro Cys Gln Glu Gly Lys Glu Tyr Thr Asp Lys Ala His Gly Leu Glu Val Glu Ile Asn Cys Thr Val Cys Glu His Cys Asp Glu Gly His Gly Leu Glu Val Glu Ile Asn Cys Thr Asp Lys Asp Glu Gly His Gly Leu Glu Val Glu Ile Asn Cys Thr Asp Eys Asp Glu His Cys Asp Glu Gly His Gly Leu Glu Val Glu Ile Asn Cys Thr Arg Thr Gln Asn Thr Lys Cys Arg Cys Asp Glu His Cys Asp Glu His Cys Asp Glu His Cys Asp Glu Gly His Gly Ile Ile Lys Glu Cys Thr Leu Thr Gla Asn Thr Lys Cys Glu His Gly Ile Ile Lys Glu Cys Thr Leu Thr Info Cys Asn Thr Lys Cys Glu Glu Glu Glu Gly Ser Arg Ser Asn Leu Gly Trp Leu Cys Leu Leu Leu Leu Pro Ile Pro Leu Ile Val Trp Val Lys Arg Lys Glu Val Glu Lys Thr Cys Arg Lys His Arg Lys Glu Asn Gln Gly

Ser His Glu Ser Pro Thr Leu Asn Pro Glu Thr 210 215

<210> 129

<211> 280

<212> PRT

<213> Rattus rattus

<400> 129

Met Gly Leu Ser Thr Val Pro Asp Leu Leu Leu Pro Leu Val Leu Leu 5 10 15

Glu Leu Leu Val Gly Ile Tyr Pro Ser Gly Val Ile Gly Leu Val Pro 20 25 30

His Leu Gly Asp Arg Glu Lys Arg Asp Ser Val Cys Pro Gln Gly Lys 35 40 45

Tyr Ile His Pro Gln Asn Asn Ser Ile Cys Cys Thr Lys Cys His Lys 50 55 60

A-378CIP5 rev.ST25.txt
Gly Thr Tyr Leu Tyr Asn Asp Cys Pro Gly Pro Gly Gln Asp Thr Asp
65 70 75 80 Cys Arg Glu Cys Glu Ser Gly Ser Phe Thr Ala Ser Glu Asn His Leu 85 90 95 Arg His Cys Leu Ser Cys Ser Lys Cys Arg Lys Glu Met Gly Gln Val 100 105 110 Glu Ile Ser Ser Cys Thr Val Asp Arg Asp Thr Val Cys Gly Cys Arg 115 120 125Lys Asn Gln Tyr Arg His Tyr Trp Ser Glu Asn Leu Phe Gln Cys Phe 130 135 140 Asn Cys Ser Leu Cys Leu Asn Gly Thr Val His Leu Ser Cys Gln Glu 145 150 155 160 Lys Gln Asn Thr Val Cys Thr Cys His Ala Gly Phe Phe Leu Arg Glu 165 170 175 Asn Glu Cys Val Ser Cys Ser Asn Cys Lys Lys Ser Leu Glu Cys Thr 180 185 190 Lys Leu Cys Leu Pro Gln Ile Glu Asn Val Lys Gly Thr Glu Asp Ser 195 200 205 Gly Thr Thr Val Leu Leu Pro Leu Val Ile Phe Phe Gly Leu Cys Leu 210 215 220 Leu Ser Leu Leu Phe Ile Gly Leu Met Thr Arg Thr Gln Arg Trp Lys 235 240 Ser Lys Leu Tyr Ser Ile Val Cys Gly Lys Ser Thr Pro Glu Lys Glu 245 250 255 Gly Glu Leu Glu Gly Thr Thr Lys Pro Leu Ala Pro Asn Pro Ser 260 265 270 Phe Ser Pro Thr Pro Gly Phe Thr 275 280 <210> 130 <211> 207 <212> **PRT** <213> Rattus rattus

<400> 130

A-378CIP5 rev.ST25.txt

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1 5 10 15 Asp Asp Val Pro Tyr Ser Ser Asn Gln Gly Lys Cys Gly Gly His Asp 20 25 30 Tyr Glu Lys Asp Gly Leu Cys Cys Ala Ser Cys His Pro Gly Phe Tyr 35 40 45 Ala Ser Arg Leu Cys Gly Pro Gly Ser Asn Thr Val Cys Ser Pro Cys 50 60 Glu Asp Gly Thr Phe Thr Ala Ser Thr Asn His Ala Pro Ala Cys Val 65 70 75 80 Ser Cys Arg Gly Pro Cys Thr Gly His Leu Ser Glu Ser Gln Pro Cys 85 90 95 Asp Arg Thr His Asp Arg Val Cys Asn Cys Ser Thr Gly Asn Tyr Cys 100 105 110 Leu Leu Lys Gly Gln Asn Gly Cys Arg Ile Cys Ala Pro Gln Thr Lys 115 120 125 Cys Pro Ala Gly Tyr Gly Val Ser Gly His Thr Arg Ala Gly Asp Thr 130 135 140 Leu Cys Glu Lys Cys Pro Pro His Thr Tyr Ser Asp Ser Leu Ser Pro 145 150 155 160 Thr Glu Arg Cys Gly Thr Ser Phe Asn Tyr Ile Ser Val Gly Phe Asn 170 175 Leu Tyr Pro Val Asn Glu Thr Ser Cys Thr Thr Ala Gly His Asn 180 185 Glu Val Ile Lys Thr Lys Glu Phe Thr Val Thr Leu Asn Tyr Thr 195 200 205

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<211> 227

<212> PRT

<213> Rattus rattus

<400> 131

Met Ala Pro Val Ala Val Trp Ala Ala Leu Ala Val Gly Leu Glu Leu 1 5 10 15 A-378CIP5 rev.ST25.txt
Trp Ala Ala Ala His Ala Leu Pro Ala Gln Val Ala Phe Thr Pro Tyr
20 25 30 Ala Pro Glu Pro Gly Ser Thr Cys Arg Leu Arg Glu Thr Thr Asp Gln 35 40 45 Thr Ala Gln Met Cys Cys Ser Lys Cys Ser Pro Gly Gln His Ala Lys 50 55 60 Val Phe Cys Thr Lys Thr Ser Asp Thr Val Cys Asp Ser Cys Glu Asp 65 70 75 80 Ser Thr Tyr Thr Gln Leu Trp Asn Trp Val Pro Glu Cys Leu Ser Cys 85 90 95 Gly Ser Arg Cys Ser Ser Asp Gln Val Glu Thr Gln Ala Cys Thr Arg 100 105 110 Glu Gln Asn Arg Ile Cys Thr Cys Arg Pro Gly Trp Tyr Cys Ala Leu 115 120 125 Ser Lys Gln Glu Gly Cys Arg Leu Cys Ala Pro Leu Arg Lys Cys Arg 130 135 140 Pro Gly Phe Gly Val Ala Arg Pro Gly Thr Glu Thr Ser Asp Val Val 145 150 155 160 Cys Lys Pro Cys Ala Pro Gly Thr Phe Ser Asn Thr Thr Ser Ser Thr 165 170 175 Asp Ile Cys Arg Pro His Gln Ile Cys Asn Val Val Ala Ile Pro Gly 180 185 Asn Ala Ser Arg Asp Ala Val Cys Thr Ser Thr Ser Pro Thr Arg Ser 195 200 205 Met Ala Pro Gly Ala Val His Leu Pro Gln Pro Val Ser Thr Arg Ser 210 215 220 Gln His Thr 225

<210> 132

<211> 197

<212> PRT

<213> Rattus rattus

<400> 132 A-378CIP5 rev.ST25.txt
Met Val Ser Leu Pro Arg Leu Cys Ala Leu Trp Gly Cys Leu Leu Thr
1 5 10 Ala Val His Leu Gly Gln Cys Val Thr Cys Ser Asp Lys Gln Tyr Leu 20 25 30 His Asp Gly Gln Cys Cys Asp Leu Cys Gln Pro Gly Ser Arg Leu Thr 35 40 45 Ser His Cys Thr Ala Leu Glu Lys Thr Gln Cys His Pro Cys Asp Ser 50 60 Gly Glu Phe Ser Ala Gln Trp Asn Arg Glu Ile Arg Cys His Gln His 65 70 75 80 Arg His Cys Glu Pro Asn Gln Gly Leu Arg Val Lys Lys Glu Gly Thr 85 90 95 Ala Glu Ser Asp Thr Val Cys Thr Cys Lys Glu Gly Gln His Cys Thr 100 105 110Ser Lys Asp Cys Glu Ala Cys Ala Gln His Thr Pro Cys Ile Pro Gly 115 120 Phe Gly Val Met Glu Met Ala Thr Glu Thr Thr Asp Thr Val Cys His 130 135 140 Pro Cys Pro Val Gly Phe Phe Ser Asn Gln Ser Ser Leu Phe Glu Lys 145 150 155 160 Cys Tyr Pro Trp Thr Ser Cys Glu Asp Lys Asn Leu Glu Val Leu Gln 165 170 175 Lys Gly Thr Ser Gln Thr Asn Val Ile Cys Gly Leu Lys Ser Arg Met 180 185 190

Arg Ala Leu Leu Val 195

<210> 133

<211> 208

<212> PRT

<213> Rattus rattus

<400> 133

Met Asn Lys Trp Leu Cys Cys Ala Leu Leu Val Phe Leu Asp Ile Ile 1 5 10 15 A-378CIP5 rev.ST25.txt Glu Trp Thr Thr Gln Glu Thr Phe Pro Pro Lys Tyr Leu His Tyr Asp 20 25 30Pro Glu Thr Gly Arg Gln Leu Leu Cys Asp Lys Cys Ala Pro Gly Thr 35 40 45 Tyr Leu Lys Gln His Cys Thr Val Arg Arg Lys Thr Leu Cys Val Pro 50 55 60 Cys Pro Asp Tyr Ser Tyr Thr Asp Ser Trp His Thr Ser Asp Glu Cys 65 70 75 80 Val Tyr Cys Ser Pro Val Cys Lys Glu Leu Gln Thr Val Lys Gln Glu 85 90 95 Cys Asn Arg Thr His Asn Arg Val Cys Glu Cys Glu Glu Gly Arg Tyr 100 105 110Leu Glu Leu Glu Phe Cys Leu Lys His Arg Ser Cys Pro Pro Gly Leu 115 120 125 Gly Val Leu Gln Ala Gly Thr Pro Glu Arg Asn Thr Val Cys Lys Arg 130 135 140 Cys Pro Asp Gly Phe Phe Ser Gly Glu Thr Ser Ser Lys Ala Pro Cys 145 150 155 160 Arg Lys His Thr Asn Cys Ser Ser Leu Gly Leu Leu Leu Ile Gln Lys 165 170 175 Gly Asn Ala Thr His Asp Asn Val Cys Ser Gly Asn Arg Glu Ala Thr 180 185 Gln Asn Cys Gly Ile Asp Val Thr Leu Cys Glu Glu Ala Phe Phe Arg 195 200 205

<210> 134

<211> 224

<212> PRT

<213> Rattus rattus

<400> 134

Met Gly Ala Gly Ala Thr Gly Arg Ala Met Asp Gly Pro Arg Leu Leu $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$

Leu Leu Leu Leu Gly Val Ser Leu Gly Gly Ala Lys Glu Ala Cys 20 25 30 A-378CIP5 rev.ST25.txt Pro Thr Gly Leu Tyr Thr His Ser Gly Glu Cys Cys Lys Ala Cys Asn 35 40 45 Leu Gly Glu Gly Val Ala Gln Pro Cys Gly Ala Asn Gln Thr Val Cys 50 60 Glu Pro Cys Leu Asp Ser Val Thr Phe Ser Asp Val Val Ser Ala Thr 65 70 75 80 Glu Pro Cys Lys Pro Cys Thr Glu Cys Val Gly Leu Gln Ser Met Ser 85 90 95 Ala Pro Cys Val Glu Ala Asp Asp Ala Val Cys Arg Cys Ala Tyr Gly 100 105 110 Tyr Tyr Gln Asp Glu Thr Thr Gly Arg Cys Glu Ala Cys Arg Val Cys 115 120 125 Glu Ala Gly Ser Gly Leu Val Phe Ser Cys Gln Asp Lys Gln Asn Thr 130 135 140 Val Cys Glu Glu Cys Pro Asp Gly Thr Tyr Ser Asp Glu Ala Asn His 145 150 155 160 Val Asp Pro Cys Leu Pro Cys Thr Val Cys Glu Asp Thr Glu Arg Gln
165 170 175 Leu Arg Glu Cys Thr Arg Trp Ala Asp Ala Glu Cys Glu Glu Ile Pro 180 185 190 Gly Arg Trp Ile Thr Arg Ser Thr Pro Pro Glu Gly Ser Asp Ser Thr 195 200 205 Ala Pro Ser Thr Gln Glu Pro Glu Ala Pro Pro Glu Gln Asp Leu Ile 210 215 220 <210> 135 <211> 202 <212> **PRT** <213> Rattus rattus

<400> 135

Met Tyr Val Trp Val Gln Gln Pro Thr Ala Phe Leu Leu Gly Leu
1 5 10 15

Ser Leu Gly Val Thr Val Lys Leu Asn Cys Val Lys Asp Thr Tyr Pro
20 25 30

Page 53

A-378CIP5 rev.ST25.txt
Ser Gly His Lys Cys Cys Arg Glu Cys Gln Pro Gly His Gly Met Val
35 40 45

Ser Arg Cys Asp His Thr Arg Asp Thr Val Cys His Pro Cys Glu Pro 50 60

Gly Phe Tyr Asn Glu Ala Val Asn Tyr Asp Thr Cys Lys Gln Cys Thr 65 70 75 80

Gln Cys Asn His Arg Ser Gly Ser Glu Leu Lys Gln Asn Cys Thr Pro 85 90 95

Thr Glu Asp Thr Val Cys Gln Cys Arg Pro Gly Thr Gln Pro Arg Gln 100 105 110

Asp Ser Ser His Lys Leu Gly Val Asp Cys Val Pro Cys Pro Pro Gly 115 120

His Phe Ser Pro Gly Ser Asn Gln Ala Cys Lys Pro Trp Thr Asn Cys 130 140

Thr Leu Ser Gly Lys Gln Ile Arg His Pro Ala Ser Asn Ser Val Cys 145 150 155 160

Glu Asp Arg Ser Leu Leu Ala Thr Leu Leu Trp Glu Thr Gln Arg Thr 165 170 175

Thr Phe Arg Pro Thr Thr Val Pro Ser Thr Thr Val Trp Pro Arg Thr 180 185 190

Ser Gln Leu Pro Ser Thr Pro Thr Leu Val 195 200

<210> 136

<211> 191

<212> PRT

<213> Rattus rattus

<400> 136

Met Gly Asn Asn Cys Tyr Asn Val Val Val Ile Val Leu Leu Val 10 15

Gly Cys Glu Lys Val Gly Ala Val Gln Asn Ser Cys Asp Asn Cys Gln
20 25 30

Pro Gly Thr Phe Cys Arg Lys Tyr Asn Pro Val Cys Lys Ser Cys Pro 35 40 45

A-378CIP5 rev.ST25.txt
Pro Ser Thr Phe Ser Ser Ile Gly Gly Gln Pro Asn Cys Asn Ile Cys
50 55 60 Arg Val Cys Ala Gly Tyr Phe Arg Phe Lys Lys Phe Cys Ser Ser Thr 65 70 75 80 His Asn Ala Glu Cys Glu Cys Ile Glu Gly Phe His Cys Leu Gly Pro 85 90 95 Gln Cys Thr Arg Cys Glu Lys Asp Cys Arg Pro Gly Gln Glu Leu Thr 100 105 110 Lys Gln Gly Cys Lys Thr Cys Ser Leu Gly Thr Phe Asn Asp Gln Asn 115 120 125 Gly Thr Gly Val Cys Arg Pro Trp Thr Asn Cys Ser Leu Asp Gly Arg 130 135 140 Ser Val Leu Lys Thr Gly Thr Thr Glu Lys Asp Val Val Cys Gly Pro 145 150 155 160 Pro Val Val Ser Phe Ser Pro Ser Thr Thr Ile Ser Val Thr Pro Glu 165 170 175 Gly Gly Pro Gly Gly His Ser Leu Gln Val Leu Thr Leu Phe Leu 180 185 190 <210> 137 <211> 54 <212> DNA <213> Artificial Sequence <220> <223> huOPG

<400> 137

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<210> 138

<211> 120

<212> PRT

<213> Homo sapiens

<400> 138

His Ala Leu Pro Ala Gln Val Ala Phe Thr Pro Tyr Ala Pro Glu Pro 1 5 10 15 Page 55 54

Gly Ser Thr Cys Arg Leu Arg Glu Tyr Tyr Asp Gln Thr Ala Gln Met 20 25 30

Cys Cys Ser Lys Cys Ser Pro Gly Gln His Ala Lys Val Phe Cys Thr 35 40 45

Lys Thr Ser Asp Thr Val Cys Asp Ser Cys Glu Asp Ser Thr Tyr Thr 50 55 60

Gln Leu Trp Asn Trp Val Pro Glu Cys Leu Ser Cys Gly Ser Arg Cys 65 70 75 80

Ser Ser Asp Gln Val Glu Thr Gln Ala Cys Thr Arg Glu Gln Asn Arg 85 90 95

Ile Cys Thr Cys Arg Pro Gly Trp Tyr Cys Ala Leu Ser Lys Gln Glu 100 105 110

Gly Cys Arg Leu Cys Ala Pro Leu 115 120

<210> 139

<211> 380

<212> PRT

<213> Homo sapiens

<400> 139

Glu Thr Phe Pro Pro Lys Tyr Leu His Tyr Asp Glu Glu Thr Ser His 1 10 15

Gln Leu Leu Cys Asp Lys Cys Pro Pro Gly Thr Tyr Leu Lys Gln His 20 25 30

Cys Thr Ala Lys Trp Lys Thr Val Cys Ala Pro Cys Pro Asp His Tyr 35 40 45

Tyr Thr Asp Ser Trp His Thr Ser Asp Glu Cys Leu Tyr Cys Ser Pro 50 55 60

Val Cys Lys Glu Leu Gln Tyr Val Lys Gln Glu Cys Asn Arg Thr His 65 70 75 80

Asn Arg Val Cys Glu Cys Lys Glu Gly Arg Tyr Leu Glu Ile Glu Phe 85 90 95

Cys Leu Lys His Arg Ser Cys Pro Pro Gly Phe Gly Val Val Gln Ala 100 105 110 Page 56

Gly Thr Pro Glu Arg Asn Thr Val Cys Lys Arg Cys Pro Asp Gly Phe 115 120 125 Phe Ser Asn Glu Thr Ser Ser Lys Ala Pro Cys Arg Lys His Thr Asn 130 140 Cys Ser Val Phe Gly Leu Leu Leu Thr Gln Lys Gly Asn Ala Thr His 145 150 155 160 Asp Asn Ile Cys Ser Gly Asn Ser Glu Ser Thr Gln Lys Cys Gly Ile 165 170 175Asp Val Thr Leu Cys Glu Glu Ala Phe Phe Arg Phe Ala Val Pro Thr 180 185 190 Lys Phe Thr Pro Asn Trp Leu Ser Val Leu Val Asp Asn Leu Pro Gly
195 200 205 Thr Lys Val Asn Ala Glu Ser Val Glu Arg Ile Lys Arg Gln His Ser 210 215 220 Ser Gln Glu Gln Thr Phe Gln Leu Leu Lys Leu Trp Lys His Gln Asn 225 230 235 240 Lys Ala Gln Asp Ile Val Lys Lys Ile Ile Gln Asp Ile Asp Leu Cys 245 250 255 Glu Asn Ser Val Gln Arg His Ile Gly His Ala Asn Leu Thr Phe Glu 260 265 270 Gln Leu Arg Ser Leu Met Glu Ser Leu Pro Gly Lys Lys Val Gly Ala 275 280 285 Glu Asp Ile Glu Lys Thr Ile Lys Ala Cys Lys Pro Ser Asp Gln Ile 290 295 300 Leu Lys Leu Leu Ser Leu Trp Arg Ile Lys Asn Gly Asp Gln Asp Thr 305 310 315 320 Leu Lys Gly Leu Met His Ala Leu Lys His Ser Lys Thr Lys His Phe 325 330 335 Pro Lys Thr Val Thr Gln Ser Leu Lys Lys Thr Ile Arg Phe Leu His 340 345 Ser Phe Thr Met Tyr Lys Leu Tyr Gln Lys Leu Phe Leu Glu Met Ile 355 360 365 Gly Asn Gln Val Gln Ser Val Lys Ile Ser Cys Leu 370 375 380 Page 57

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y		
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24.0		
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<223>	huOPG	
<400> gtttct	143 tcgt cataatgaag ctgggtggtc c	31
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	144 cacc caggacgaag aaacctctc	29
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<400> gagagg	145 tttc ttcgtcctgg gtggtccac	29
<210>	146	
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<210>	148	
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<210>	149	
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<210>	151	
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24

tgacctctcg gaaagcagcg tgca

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tycacy	ctgc tttccgagag gtca			24
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<400>	156 aatc gagcgagcag ctcc			24
ceeega	aace gagegageag ecce			- '
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Asp Ser Trp His Thr Ser 65 70

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Tyr Leu Lys Gln His Cys Thr Val Arg Arg Lys Thr Leu Cys Val Pro 50 55 60

Cys Pro Asp His Ser Tyr Thr Asp Ser Trp His Thr Ser Asp Glu Cys 65 70 75 80 Page 66

Val Tyr Cys Ser Pro Val Cys Lys Glu Leu Gln Ser Val Lys Gln Glu 85 90 95 Cys Asn Arg Thr His Asn Arg Val Cys Glu Cys Glu Glu Gly Arg Tyr 100 105 110 Leu Glu Ile Glu Phe Cys Leu Lys His Arg Ser Cys Pro Pro Gly Ser 115 120 125 Gly Val Val Gln Ala Gly Thr Pro Glu Arg Asn Thr Val Cys Lys 130 135 140 Cys Pro Asp Gly Phe Phe Ser Gly Glu Thr Ser Ser Lys Ala Pro Cys 145 150 155 160 Ile Lys His Thr Asn Cys Ser Thr Phe Gly Leu Leu Leu Ile Gln Lys 165 170 175 Gly Asn Ala Thr His Asp Asn Val Cys Ser Gly Asn Arg Glu Ala Thr 180 185 190 Gln Lys Cys Gly Ile Asp Val Thr Leu Cys Glu Glu Ala Phe Phe Arg 195 200 205 Phe Ala Val Pro Thr Lys Ile Ile Pro Asn Trp Leu Ser Val Leu Val 210 215 220 Asp Ser Leu Pro Gly Thr Lys Val Asn Ala Glu Ser Val Glu Arg Ile 225 230 235 240 Lys Arg Arg His Ser Ser Gln Glu Gln Thr Phe Gln Leu Leu Lys Leu 245 250 255 Trp Lys His Gln Asn Arg Asp Gln Glu Met Val Lys Lys Ile Ile Gln
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Lys Thr Ser His Phe Pro Lys Thr Val Thr His Ser Leu Arg Lys Thr 355 360 365

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Val Tyr Cys Ser Pro Val Cys Lys Glu Leu Gln Thr Val Lys Gln Glu 85 90 95

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Cys Pro Asp Gly Phe Phe Ser Gly Glu Thr Ser Ser Lys Ala Pro Cys 145 150 155 160 Page 68

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Cys Asn Arg Thr His Asn Arg Val Cys Glu Cys Lys Glu Gly Arg Tyr 100 105 110

Leu Glu Ile Glu Phe Cys Leu Lys His Arg Ser Cys Pro Pro Gly Leu 115 120 125

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Cys Pro Asp Gly Phe Phe Ser Asn Glu Thr Ser Ser Lys Ala Pro Cys 155 160

Arg Lys His Thr Asn Cys Ser Val Phe Gly Leu Leu Leu Thr Gln Lys 165 170 175

Gly Asn Ala Thr His Asp Asn Ile Cys Ser Gly Asn Ser Glu Ser Thr 180 185 190

Gln Lys Cys Gly Ile Asp Val Thr Leu Cys Glu Glu Ala Phe Phe Arg 195 200 205

Phe Ala Val Pro Thr Lys Phe Thr Pro Asn Trp Leu Ser Val Leu Val 210 220

Asp Asn Leu Pro Gly Thr Lys Val Asn Ala Glu Ser Val Glu Arg Ile 235 240 Page 70

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Lys Trp Lys Thr Val Cys Ala Pro Cys Pro Asp His Tyr Tyr Thr Asp 35 40 45 Page 71

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Gly Thr Pro Glu Arg Asn Thr Val Cys Lys Lys Cys Pro Asp Gly Phe 115 120 125

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gag Glu	gag Glu	cag Gln 270	tac Tyr	aac Asn	agc Ser	acg Thr	tac Tyr 275	cgt Arg	gtg Val	gtc Val	agc Ser	gtc Val 280	ctc Leu	acc Thr	gtc Val	5608
ctg Leu	cac His 285	cag Gln	gac Asp	tgg Trp	ctg Leu	aat Asn 290	ggc Gly	aag Lys	gag Glu	tac Tyr	aag Lys 295	tgc Cys	aag Lys	gtc Val	tcc Ser	5656
								gag Glu								5704
ggg Gly	cag Gln	ccc Pro	cga Arg	gaa Glu 320	cca Pro	cag Gln	gtg Val	tac Tyr	acc Thr 325	ctg Leu	ccc Pro	cca Pro	tcc Ser	cgg Arg 330	gat Asp	5752
gag Glu	ctg Leu	acc Thr	aag Lys 335	aac Asn	cag Gln	gtc Val	agc Ser	ctg Leu 340	acc Thr	tgc Cys	ctg Leu	gtc Val	aaa Lys 345	ggc Gly	ttc Phe	5800
tat Tyr	ccc Pro	agc ser 350	gac Asp	atc Ile	gcc Ala	gtg Val	gag Glu 355	tgg Trp	gag Glu	agc Ser	aat Asn	ggg Gly 360	cag Gln	ccg Pro	gag Glu	5848
								gtg Val	Leu		Ser 375					5896

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ttc ctc Phe Leu 380	tac agc Tyr Ser	aag ctc Lys Leu 385	acc gtg Thr Val	gac a Asp L	aag agc _ys Ser 390	agg tgg Arg Trp	cag Gln	cag Gln	ggg G1y 395	5944
aac gtc Asn Val	ttc tca Phe Ser	tgc tcc Cys Ser 400	gtg atg Val Met	cat g His G	gag gct Glu Ala 405	ctg cac Leu His	aac Asn	cac His 410	tac Tyr	5992
acg cag Thr Gln	aag agc Lys Ser 415	ctc tcc Leu Ser	ctg tct Leu Ser	ccg g Pro G 420	ggt aaa Gly Lys	tgataac	tcg a	ac		6037

<210> 177

<211> 422

<212> PRT

<213> Homo sapiens

<400> 177

Met Asn Lys Leu Leu Cys Cys Ala Leu Val Phe Leu Asp Ile Ser Ile 1 5 10 15

Lys Trp Thr Thr Gln Glu Thr Phe Pro Pro Lys Tyr Leu His Tyr Asp $20 \hspace{1cm} 25 \hspace{1cm} 30$

Glu Glu Thr Ser His Gln Leu Leu Cys Asp Lys Cys Pro Pro Gly Thr 35 40 45

Tyr Leu Lys Gln His Cys Thr Ala Lys Trp Lys Thr Val Cys Ala Pro 50 55 60

Cys Pro Asp His Tyr Tyr Thr Asp Ser Trp His Thr Ser Asp Glu Cys 65 70 75 80

Leu Tyr Cys Ser Pro Val Cys Lys Glu Leu Gln Tyr Val Lys Gln Glu 85 90 95

Cys Asn Arg Thr His Asn Arg Val Cys Glu Cys Lys Glu Gly Arg Tyr 100 105 110

Leu Glu Ile Glu Phe Cys Leu Lys His Arg Ser Cys Pro Pro Gly Phe 115 120 125

Gly Val Val Gln Ala Glỳ Thr Pro Glu Arg Asn Thr Val Cys Lys Arg 130 135 140

Cys Pro Asp Gly Phe Phe Ser Asn Glu Thr Ser Ser Lys Ala Pro Cys 145 150 155 160

Arg Lys His Thr Asn Cys Ser Val Phe Gly Leu Leu Leu Thr Gln Lys 165 170 175 Page 78

Gly Asn Ala Thr His Asp Asn Ile Cys Ser Gly Asn Ser Glu Ser Thr 180 185 190 Gln Lys Val Asp Lys Thr His Thr Cys Pro Pro Cys Pro Ala Pro Glu 195 200 205 Leu Leu Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp 210 215 220 Thr Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val Asp 225 230 235 240 Val Ser His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr Val Asp Gly 245 250 255 Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln Tyr Asn 260 265 270 Ser Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln Asp Trp 275 280 285 Leu Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Ala Leu Pro 290 295 300 Ala Pro Ile Glu Lys Thr Ile Ser Lys Ala Lys Gly Gln Pro Arg Glu 305 310 315 320 Pro Gln Val Tyr Thr Leu Pro Pro Ser Arg Asp Glu Leu Thr Lys Asn 325 330 335 Gln Val Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile 340 345 350 Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr 355 360 365 Thr Pro Pro Val Leu Asp Ser Asp Gly Ser Phe Phe Leu Tyr Ser Lys 370 380 Leu Thr Val Asp Lys Ser Arg Trp Gln Gln Gly Asn Val Phe Ser Cys 385 390 395 400 Ser Val Met His Glu Ala Leu His Asn His Tyr Thr Gln Lys Ser Leu 405 410 415 Ser Leu Ser Pro Gly Lys 420

<210> 178



<211> 51

<212> PRT

<213> Homo sapiens

<400> 178

Tyr His Tyr Tyr Asp Gln Asn Gly Arg Met Cys Glu Glu Cys His Met $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$

Cys Gln Pro Gly His Phe Leu Val Lys His Cys Lys Gln Pro Lys Arg 20 25 30

Asp Thr Val Cys His Lys Pro Cys Glu Pro Gly Val Thr Tyr Thr Asp 35 40 45

Asp Trp His 50

CM